

## SEQUENCE LISTING

## (1) GENERAL INFORMATION:

## (i) APPLICANT:

- (A) NAME: BASF Aktiengesellschaft
- (B) STREET: Carl-Bosch-Strasse 38
- (C) CITY: Ludwigshafen
- (E) COUNTRY: Federal Republic of Germany
- (F) POSTAL CODE: D-67056
- (G) TELEPHONE: 0621/6048526
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- (I) TELEX: 1762175170

(ii) TITLE OF APPLICATION: Method for diagnosing disorders by analysis of genes

(iii) NUMBER OF SEQUENCES: 2

## (iv) COMPUTER READABLE FORM:

- (A) MEDIUM TYPE: Floppy disk
- (B) COMPUTER: IBM PC compatible
- (C) OPERATING SYSTEM: PC-DOS/MS-DOS
- (D) SOFTWARE: PatentIn Release #1.0, Version #1.25 (EPA)

## (2) INFORMATION FOR SEQ ID NO: 1:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 1517 base pairs
- (B) TYPE: Nucleic acid
- (C) STRANDEDNESS: Double
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA for mRNA

(iii) HYPOTHETICAL: NO

(iv) ANTISENSE: NO

(vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo sapiens

(ix) FEATURES:

- (A) NAME/KEY: CDS

- (B) LOCATION: 1..1024

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 1:

ATG	GGG	GAG	ATG	GAG	CAA	CTG	CGT	CAG	GAA	GCG	GAG	CAG	CTC	AAG	AAG	48
Met	Gly	Glu	Met	Glu	Gln	Leu	Arg	Gln	Glu	Ala	Glu	Gln	Leu	Lys	Lys	
1				5				10					15			
CAG	ATT	GCA	GAT	GCC	AGG	AAA	GCC	TGT	GCT	GAC	GTT	ACT	CTG	GCA	GAG	96
Gln	Ile	Ala	Asp	Ala	Arg	Lys	Ala	Cys	Ala	Asp	Val	Thr	Leu	Ala	Glu	
			20				25					30				
CTG	GTG	TCT	GGC	CTA	GAG	GTG	GTG	GGA	CGA	GTC	CAG	ATG	CGG	ACG	CGG	144
Leu	Val	Ser	Gly	Leu	Glu	Val	Val	Gly	Arg	Val	Gln	Met	Arg	Thr	Arg	
		35				40					45					
CGG	ACG	TTA	AGG	GGA	CAC	CTG	GCC	AAG	ATT	TAC	GCC	ATG	CAC	TGG	GCC	192
Arg	Thr	Leu	Arg	Gly	His	Leu	Ala	Lys	Ile	Tyr	Ala	Met	His	Trp	Ala	
	50				55			60								
ACT	GAT	TCT	AAG	CTG	CTG	GTA	AGT	GCC	TCG	CAA	GAT	GGG	AAG	CTG	ATC	240
Thr	Asp	Ser	Lys	Leu	Leu	Val	Ser	Ala	Ser	Gln	Asp	Gly	Lys	Leu	Ile	
	65			70				75					80			
GTG	TGG	GAC	AGC	TAC	ACC	ACC	AAC	AAG	GTG	CAC	GCC	ATC	CCA	GTG	CGC	288
Val	Trp	Asp	Ser	Tyr	Thr	Thr	Asn	Lys	Val	His	Ala	Ile	Pro	Leu	Arg	
			85				90					95				
TCC	TCC	TGG	GTC	ATG	ACC	TGT	GCC	TAT	GCC	CCA	TCA	GGG	AAC	TTT	GTG	336
Ser	Ser	Trp	Val	Met	Thr	Cys	Ala	Tyr	Ala	Pro	Ser	Gly	Asn	Phe	Val	

					100					105					110					
GCA	TGT	GGG	GGG	CTG	GAC	AAC	ATG	TCC	ATC	TAC	AAC	CTC	AAA	TCC	384					
Ala	Cys	Gly	Gly	Leu	Asp	Asn	Met	Cys	Ser	Ile	Tyr	Asn	Leu	Lys	Ser					
		115					120					125								
CGT	GAG	GGC	AAT	GTC	AAG	GTC	AGC	CGG	GAG	CTT	TCT	GCT	CAC	ACA	GGT					
Arg	Glu	Gly	Asn	Val	Lys	Val	Ser	Arg	Glu	Leu	Ser	Ala	His	Thr	Gly					
		130					135					140								
TAT	CTC	TCC	TGC	TGC	CGC	TTC	CTG	GAT	GAC	AAC	AAT	GTG	ACC	AGC						
Tyr	Leu	Ser	Cys	Cys	Arg	Phe	Leu	Asp	Asp	Asn	Asn	Ile	Val	Thr	Ser					
		145					150					155			160					
TCG	GGG	GAC	ACC	ACG	TGT	GCC	TTG	TGG	GAC	ATT	GAG	ACT	GGG	CAG	CAG					
Ser	Gly	Asp	Thr	Thr	Cys	Ala	Leu	Trp	Asp	Ile	Glu	Thr	Gly	Gln	Gln					
				165					170					175						
AAG	ACT	GTA	TTT	GTG	GGA	CAC	ACG	GGT	GAC	TGC	ATG	AGC	CTG	GCT	GTG					
Lys	Thr	Val	Phe	Val	Gly	His	Thr	Gly	Asp	Cys	Met	Ser	Leu	Ala	Val					
				180					185					190						
TCT	CCT	GAC	TTC	AAT	CTC	TTC	ATT	TCG	GGG	GCC	TGT	GAT	GCC	AGT	GCC					
Ser	Pro	Asp	Phe	Asn	Leu	Phe	Ile	Ser	Gly	Ala	Cys	Asp	Ala	Ser	Ala					
				195					200					205						
AAG	CTC	TGG	GAT	GTG	CGA	GAG	GGG	ACC	TGC	CGT	CAG	ACT	TTC	ACT	GGC					
Lys	Leu	Trp	Asp	Val	Arg	Glu	Gly	Thr	Cys	Arg	Gln	Thr	Phe	Thr	Gly					
				210					215					220						
CAC	GAG	TCG	GAC	ATC	AAC	GCC	ATC	TGT	TTC	TTC	CCC	AAT	GGA	GAG	GCC					
His	Glu	Ser	Asp	Ile	Asn	Ala	Ile	Cys	Phe	Phe	Pro	Asn	Gly	Glu	Ala					
				225							235									
ATC	TGC	ACG	GGC	TCG	GAT	GAC	GCT	TCC	TGC	CGC	TTG	TTT	GAC	CTG	CGG					
Ile	Cys	Thr	Gly	Ser	Asp	Asp	Ala	Ser	Cys	Arg	Leu	Phe	Asp	Leu	Arg					
				245							250									
GCA	GAC	CAG	GAG	CTG	ATC	TGC	TTC	TCC	CAC	GAG	AGC	ATC	ATC	TGC	GGC					
Ala	Asp	Gln	Glu	Leu	Ile	Cys	Phe	Ser	His	Glu	Ser	Ile	Ile	Cys	Gly					
				260							265									
ATC	ACG	TCT	GTG	GCC	TTC	TCC	CTC	AGT	GGC	CGC	CTA	CTA	TTC	GCT	GGC					
Ile	Thr	Ser	Val	Ala	Phe	Ser	Leu	Ser	Gly	Arg	Leu	Leu	Phe	Ala	Gly					
				275											285					
TAC	GAC	GAC	TTC	AAC	TGC	AAT	GTC	TGG	GAC	TCC	ATG	AAG	TCT	GAG	CGT					
Tyr	Asp	Asp	Phe	Asn	Cys	Asn	Val	Trp	Asp	Ser	Met	Lys	Ser	Glu	Arg					
				290											300					
GTG	GGC	ATC	CTC	TCT	GGC	CAC	GAT	AAC	AGG	GTG	AGC	TGC	CTG	GGA	GTC					
Val	Gly	Ile	Leu	Ser	Gly	His	Asp	Asn	Arg	Val	Ser	Cys	Leu	Gly	Val					
				305											310					
ACA	GCT	GAC	GGG	ATG	GCT	GTG	GCC	ACA	GGT	TCC	TGG	GAC	AGC	TTC	CTC					
Thr	Ala	Asp	Gly	Met	Ala	Val	Ala	Thr	Gly	Ser	Trp	Asp	Ser	Phe	Leu					
				325											335					
AAA	ATC	TGG	AAC	TGA	G	GAGGCTGGAG	AAAGGGAAGT	GGAAGGCAGT	GAACACACTC	1064										
Lys	Ile	Trp	Asn	*																
				340																
AGCAGCCCCC	TGCCCCGACCC	CATCTCATTC	AGGTGTTCTC	TTCTATATTC	CGGGTGCCAT	1124														
TCCCACTAAG	CTTTCTCCTT	TGAGGGCAGT	GGGGAGCATG	GGACTGTGCC	TTTGGGAGGC	1184														
AGCATCAGGG	ACACAGGGGC	AAAGAAGTGC	CCCATCTCCT	CCCATGGCCT	TCCCTCCCCA	1244														
CAGTCCTCAC	AGCCTCTCCC	TTAATGAGCA	AGGACAACCT	GCCCCTCCCC	AGCCCTTTGC	1304														
AGGCCCAGCA	GACTTGAGTC	TGAGGCCCCA	GGCCCTAGGA	TTCTCTCCCC	AGAGCCACTA	1364														
CCTTTGTCCA	GGCCTGGGTG	GTATAGGGCG	TTTGGCCCTG	TGACTATGGC	TCTGGCACCA	1424														
CTAGGGTCCT	GGCCCTCTTC	TTATTCATGC	TTTCTCCTTT	TTCTACCTTT	TTTTCTCTCC	1484														
TAAGACACCT	GCAATAAAGT	GTAGCACCCCT	GGT			1517														

## (2) INFORMATION FOR SEQ ID NO: 2:

## (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 341 amino acids

(B) TYPE: Amino acid

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: Protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 2:

Met	Gly	Glu	Met	Glu	Gln	Leu	Arg	Gln	Glu	Ala	Glu	Gln	Leu	Lys	Lys	1	5	10	15
Gln	Ile	Ala	Asp	Ala	Arg	Lys	Ala	Cys	Ala	Asp	Val	Thr	Leu	Ala	Glu	20	25	30	
Leu	Val	Ser	Gly	Leu	Glu	Val	Val	Gly	Arg	Val	Gln	Met	Arg	Thr	Arg	35	40	45	
Arg	Thr	Leu	Arg	Gly	His	Leu	Ala	Lys	Ile	Tyr	Ala	Met	His	Trp	Ala	50	55	60	
Thr	Asp	Ser	Lys	Leu	Leu	Val	Ser	Ala	Ser	Gln	Asp	Gly	Lys	Leu	Ile	65	70	75	80
Val	Trp	Asp	Ser	Tyr	Thr	Thr	Asn	Lys	Val	His	Ala	Ile	Pro	Leu	Arg	85	90	95	
Ser	Ser	Trp	Val	Met	Thr	Cys	Ala	Tyr	Ala	Pro	Ser	Gly	Asn	Phe	Val	100	105	110	
Ala	Cys	Gly	Gly	Leu	Asp	Asn	Met	Cys	Ser	Ile	Tyr	Asn	Leu	Lys	Ser	115	120	125	
Arg	Glu	Gly	Asn	Val	Lys	Val	Ser	Arg	Glu	Leu	Ser	Ala	His	Thr	Gly	130	135	140	
Tyr	Leu	Ser	Cys	Cys	Arg	Phe	Leu	Asp	Asp	Asn	Asn	Ile	Val	Thr	Ser	145	150	155	160
Ser	Gly	Asp	Thr	Thr	Cys	Ala	Leu	Trp	Asp	Ile	Glu	Thr	Gly	Gln	Gln	165	170	175	
Lys	Thr	Val	Phe	Val	Gly	His	Thr	Gly	Asp	Cys	Met	Ser	Leu	Ala	Val	180	185	190	
Ser	Pro	Asp	Phe	Asn	Leu	Phe	Ile	Ser	Gly	Ala	Cys	Asp	Ala	Ser	Ala	195	200	205	
Lys	Leu	Trp	Asp	Val	Arg	Glu	Gly	Thr	Cys	Arg	Gln	Thr	Phe	Thr	Gly	210	215	220	
His	Glu	Ser	Asp	Ile	Asn	Ala	Ile	Cys	Phe	Phe	Pro	Asn	Gly	Glu	Ala	225	230	235	240
Ile	Cys	Thr	Gly	Ser	Asp	Asp	Ala	Ser	Cys	Arg	Leu	Phe	Asp	Leu	Arg	245	250	255	
Ala	Asp	Gln	Glu	Leu	Ile	Cys	Phe	Ser	His	Glu	Ser	Ile	Ile	Cys	Gly	260	265	270	
Ile	Thr	Ser	Val	Ala	Phe	Ser	Leu	Ser	Gly	Arg	Leu	Leu	Phe	Ala	Gly	275	280	285	
Tyr	Asp	Asp	Phe	Asn	Cys	Asn	Val	Trp	Asp	Ser	Met	Lys	Ser	Glu	Arg	290	295	300	
Val	Gly	Ile	Leu	Ser	Gly	His	Asp	Asn	Arg	Val	Ser	Cys	Leu	Gly	Val	305	310	315	320
Thr	Ala	Asp	Gly	Met	Ala	Val	Ala	Thr	Gly	Ser	Trp	Asp	Ser	Phe	Leu	325	330	335	
Lys	Ile	Trp	Asn	*												340			

We claim:

1. The use of a genetic modification in the gene for human G protein  $\beta 3$  subunit for the diagnosis of diseases.
2. The use of a genetic modification in the gene for human G protein  $\beta 3$  subunit for establishing the risk of developing a disorder associated with G protein dysregulation.
3. The use as claimed in claim 2, wherein the genetic modification is in the codon for amino acid 275 in SEQ ID NO:1.
4. The use as claimed in claim 3, wherein there is substitution of cytosine by thymine in position 825 in SEQ ID NO:1.
5. The use as claimed in claim 2, wherein the disorder is a cardiovascular disease, a metabolic disturbance or an immunological disease.
6. The use as claimed in claim 2, wherein the disorder is hypertension.
7. A method for establishing a relative risk of developing disorders associated with G protein dysregulation for a subject, which comprises comparing the gene sequence for human G protein  $\beta 3$  subunit of the subject with the gene sequence SEQ ID NO:1, and, in the event that a thymine (T) is present at position 825, assigning the subject an increased risk of disease.
8. A method as claimed in claim 7, wherein the comparison of genes is carried out by sequencing.
9. A method as claimed in claim 8, wherein a gene section which includes position 825 is amplified before the sequencing.
10. A method as claimed in claim 7, wherein the comparison of genes is carried out by hybridization.
11. A method as claimed in claim 7, wherein the comparison of genes is carried out by cleavage using restriction enzymes.
12. A method as claimed in claim 11, wherein the restriction enzyme Dsa I is used.

The use of a genetic modification in the gene for human G protein  $\beta 3$  subunit for the diagnosis of diseases

## 5 Abstract

The present invention relates to the use of a genetic modification in the gene for human G protein  $\beta 3$  subunit for the diagnosis of diseases.

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